Serial No.: 09/423,715

Filed: January 12, 2000

Page: 2

In the Claims

Please amend the claims by replacing all prior versions of the claims pursuant to 37 C.F.R. §1.121 as modified by 68 Fed. Reg. 38611 (June 30, 2003) as follows:

 (Currently Amended) A topical composition for mutual enhancement of transdermal permeation of at least a first and a second pharmaceutically acceptable components which are both pharmacologically active agents, the composition comprising

an emulsion of at least one discontinuous phase in a continuous phase, the or each discontinuous phase comprising a eutectic mixture of first and second pharmacologically active agents and the continuous phase comprising a pharmaceutically acceptable carrier, the eutectic mixture having a melting point below 40°C; and

at least one compatible emulsifying agent,

wherein when the first pharmacologically active agent is a local anesthetic, the second pharmacologically active agent is not a local anaesthetic, wherein when the second pharmacologically active agent is a local anesthetic, the first pharmacologically active agent is not a local anesthetic, and wherein the first and the second pharmacologically active agents are each a prophylactic or a therapeutic agent, and

wherein one of the first or the second pharmacologically chlorocresol, active agent is chlorbutanol, methyl nicotinate, triprolidine, promethazine, trimeprazine, sulfiram, oxybutynin, testosterone enanthate or choline salicylate.

2. (Previously presented) The topical composition according to Claim 1, in which the first pharmacologically active agent



Serial No. : 09/423,715

Filed: January 12, 2000

Page: 3

has a melting point between 35 and 75°C, and the second pharmacologically active agent has a melting point between -40°C and 150°C.

3. (Previously presented) The topical composition according to Claim 1, in which the topical composition additionally includes, in the eutectic mixture, a third pharmaceutically acceptable component.

- 4. (Previously presented) The topical composition according to Claim 3, in which the third pharmaceutically acceptable component has a melting point between 40 and 150°C.
- 5. (Previously presented) The topical composition according to Claim 3 or 4, in which the third component is a third pharmacologically active agent.
- 6. (Previously presented) The topical composition according to Claim 3, in which the topical composition additionally includes, in the eutectic mixture, a fourth pharmaceutically acceptable component.
- 7. (Previously presented) The topical composition according to Claim 6, in which the fourth pharmaceutically acceptable component has a melting point between 40 and 150°C.
- 8. (Previously presented) The topical composition according to Claim 6 or 7, in which the fourth component comprises a fourth pharmacologically active agent.
- 9. (Previously presented) The topical composition according to Claim 1, in which said at least one discontinuous phase consists essentially of the eutectic mixture.

Pl wit

Serial No. : 09/423,715

Filed: January 12, 2000

Page: 4

10. (Canceled).

11. (Previously presented) The topical composition according to Claim 1, in which the second pharmacologically active agent is selected from the group consisting of non-steroid anti-inflammatory arylpropionic agents, narcotic analgesics, anti-fungal agents, antibacterial agents, anticholinergics, anthelmintics, antihistaminics, and antihypertensives.

12. (Previously presented) The topical composition according to Claim 8, in which the third and fourth pharmacologically active agents are each selected from the group consisting of non-steroid anti-inflammatory agents, narcotic analgesics, anti-fungal agents, antibacterial agents, anticholinergics, antihypertensives, antihistaminics, and anthelmintics.

Claim 3 or 4, in which the third pharmaceutically acceptable component is lauric acid, stearyl alcohol, menthol, thymol, cinnamic acid or an ester thereof.

- 14. (Previously presented) The topical composition according to Claim 1, in which the pharmaceutically acceptable carrier is substantially hydrophilic, said carrier comprising substantially water as the continuous phase.
- 15. (Previously presented) The topical composition according to Claim 1, in which the pharmaceutically acceptable carrier contains at least one gelling or suspension agent.
- 16. (Previously presented) The topical composition according to Claim 15, in which the gelling or suspension agent is

D'

Serial No.: 09/423,715

Filed: January 12, 2000

Page: 5

selected from the group consisting of carbomers, modified celluloses, naturally-occurring synthetic or semi-synthetic gums, modified starches, co-polymers formed between maleic anhydride and methyl vinyl ether, colloidal silica and methacrylates or a mixture thereof.

17. (Previously presented) The topical composition according to Claim 1, in which the topical composition is in the form of a gel, lotion, suspension, cream, aerosol spray, transdermal patch, medicated dressing or soft gelatin capsule.

- 18. (Previously presented) The topical composition according to Claim 1, in which the emulsifying agent is selected from the group consisting of non-ionic, cationic and anionic surfactants.
- 19. (Previously presented) The topical composition according to Claim 18, in which the emulsifying agent is a non-ionic surfactant.
- 20. (Previously presented) The topical composition according to Claim 1, in which the at least two pharmacologically active agents are structurally and/or pharmacologically diverse.

21 - 22. (Canceled)

23. (Currently Amended) A method for mutual enhancement of dermal permeation from an accessible body surface of a human of at least a first and a second pharmaceutically acceptable components which are both pharmacologically active agents, the method comprising a home of applying a topical composition for mutual enhancement

D'ut

Serial No.: 09/423,715

agent,

Filed: January 12, 2000

Page: 6

of transdermal permeation of at least first and second pharmacologically active agents, the composition comprising an emulsion of at least one discontinuous phase in a continuous phase, the or each discontinuous phase comprising a eutectic mixture of first and second pharmacologically active agents and the continuous phase comprising a pharmaceutically acceptable carrier, the eutectic mixture having a melting point below 40°C; and at least one compatible emulsifying

wherein when the first pharmacologically active local anesthetic, the pharmacologically agent is not a local anesthetic, wherein when the second pharmacologically active agent is a local anesthetic, the first pharmacologically active agent is not a local anesthetic, and wherein the first and the second pharmacologically active agents are each a prophylactic or a therapeutic agent, wherein one of the first or active pharmacologically is triclosan, agent chlorbutanol, methyl nicotinate, chlorocresol, triprolidine, promethazine, trimeprazine, sulfiram, oxybutynin, capsaicin, testosterone enanthate or choline salicylate, to-an accessible body surface of Diphereun the topical application to the human. a human in need throf is to arm

- 24. (Canceled).
- 25. (Previously presented) The topical composition according to claim 2, wherein the first pharmacologically active agent has a melting point between 40 and 50°C, and the second pharmacologically active agent has a melting point between -5 and 90°C.

P. H

Serial No. : 09/423,715

Filed : January 12, 2000

Page: 7

26. (Previously presented) The topical composition according to claim 4, wherein the third pharmaceutically acceptable component has a melting point between 40 and 75°C.

- 27. (Previously presented) The topical composition according to claim 7, wherein the fourth pharmaceutically acceptable component has a melting point between 40 and 75°C.
- 28. (Previously presented) The topical composition according to claim 11, wherein the second pharmacologically active agent is selected from the group consisting of triclosan, chlorocresol, capsaicin, trimeprazine, choline salicylate, methyl nicotinate, ibuprofen, ketoprofen, fenoprofen, flurbiprofen, etodolac, fentanyl, econazole, ketoconazole, mupirocin, chlorbutanol, clindamycin, iodine, oxybutynin, tetramisole, triprolidine, promethazine, and propranolol.
- 29. (Previously presented) The topical composition according to Claim 12, wherein the third and fourth pharmacologically active agents are each selected from the group consisting of triclosan, chlorocresol, capsaicin, trimeprazine, choline salicylate, methyl nicotinate, ibuprofen, ketoprofen, fenoprofen, flurbiprofen, etodolac, fentanyl, ketoconazole, mupirocin, econazole, chlorbutanol, clindamycin, iodine, oxybutynin, propranolol, triprolidine, promethazine, and tetramisole.
- 30. (Previously presented) The topical composition according to Claim 16, wherein the gelling or suspension agent is selected from the group consisting of xanthan gum, acacia, tragacanth, and a mixture thereof.

D'A

Serial No.: 09/423,715

Filed: January 12, 2000

Page: 8

31. (Previously presented) The topical composition according to Claim 9, in which said at least one discontinuous phase 7 consists of the eutectic mixture.

32. (Previously presented) The topical composition according to Claim 14, in which the pharmaceutically acceptable carrier is substantially hydrophilic, said carrier essentially consisting of water as the continuous phase.

33. (Previously presented) The method of claim 23, wherein the animal is a human.

- 34. (Previously presented) The method according to Claim 23, in which said at least one discontinuous phase consists essentially of the eutectic mixture.
- 35. (Previously presented) The method according to Claim 34, in which said at least one discontinuous phase consists of the eutectic mixture.
- 36. (Previously presented) The method according to Claim 23, in which the pharmaceutically acceptable carrier is substantially hydrophilic, said carrier comprising substantially water as the continuous phase.
 - 37. (Previously presented) The method according to Claim 36, in which the pharmaceutically acceptable carrier is substantially hydrophilic, said carrier essentially consisting of water as the continuous phase.

D. its